



CASE REPORT

A case of disseminated tuberculosis with increased tumour markers and testicular involvement

Metin Ozkan ^{a,*}, Canturk Tasci ^a, Arzu Balkan ^a, Ergun Ucar ^a,
Selahattin Bedir ^b, Hayati Bilgic ^a, Kudret Ekiz ^a

^a Department of Pulmonary Medicine, Gulhane Military Medical Academy, Gogus Hastaliklari Klinigi, 06018 Etlik, Ankara, Turkey

^b Department of Urology, Gulhane Military Medical Academy, 06018, Ankara, Turkey

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KEYWORDS

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Summary

We report a patient having disseminated tuberculosis (TB) with negative smear and culture for *Mycobacterium tuberculosis*, left testicular mass, and an increase in tumour markers; alpha-fetoprotein (AFP), CA-125, and CA-19-9. The patient was a 53-year-old Caucasian male, presented with night sweat, weakness, and weight loss. Radiological findings were compatible with tuberculosis, however, sputum, bronchoalveolar lavage and biopsy materials were negative for malignancy or tuberculosis. A testicular mass was detected during the physical examination and orchidectomy material revealed acid-fast bacilli containing caseating granuloma. The patient responded well to anti-tuberculous therapy; clinical and radiological improvements observed at the end of the treatment period. Tumour markers were also decreased. Crown Copyright © 2008 Published by Elsevier Ltd. All rights reserved.

Description of case

A 53-year-old male was presented with a 1-month history of night sweat, weakness, and 2-months' history of weight loss (15 kg). He was slim, pale and weak, and his body temperature was 37.8 °C on the day of admission. He was

a heavy smoker and reported using approximately 20 cigarettes per day for the previous 40 years.

A 1 × 2 cm. left testicular mass was detected by physical examination, which was also confirmed with ultrasonographic examination. No abnormality was found during the bronchoscopic examination.

White blood cells were found to be increased (12,300/μL) in complete blood count. Erythrocyte sedimentation rate was 23 mm/h, and LDH level was 298 U/L. Urinalysis was normal. HBsAg, HCV and HIV antibodies were negative. Alpha-fetoprotein (AFP) was marginally elevated (9.6 ng/mL). CA-125 (412.7 U/mL), and CA-19-9

* Corresponding author. Tel.: +90 312 304 4403; fax: +90 312 304 2010.

E-mail address: metin58@yahoo.com (M. Ozkan).

Table 1 Tumour markers before and after anti-tuberculous therapy

Marker	Before anti-tuberculous therapy	After 2 months of therapy	After 6 months of therapy
CA-125 (U/mL)(Normal: <35)	412.7	89.7	6.4
CA-19-9 (U/mL)(Normal: <37)	93.33	35.4	14.45
AFP (ng/mL)(Normal: <8.1)	9.6	8.6	7.6

(93.33 U/mL) levels were obviously high (Table 1). Sputum, bronchoscopic lavage, and transbronchial biopsy materials were negative both for microorganisms and malignancy.

Bilaterally diffuse reticulo-nodular infiltrations were observed in posteroanterior chest radiograms (Fig. 1). Abdominal computed tomography (CT) indicated para-vertebral, paraaortic, and abdominal multiple

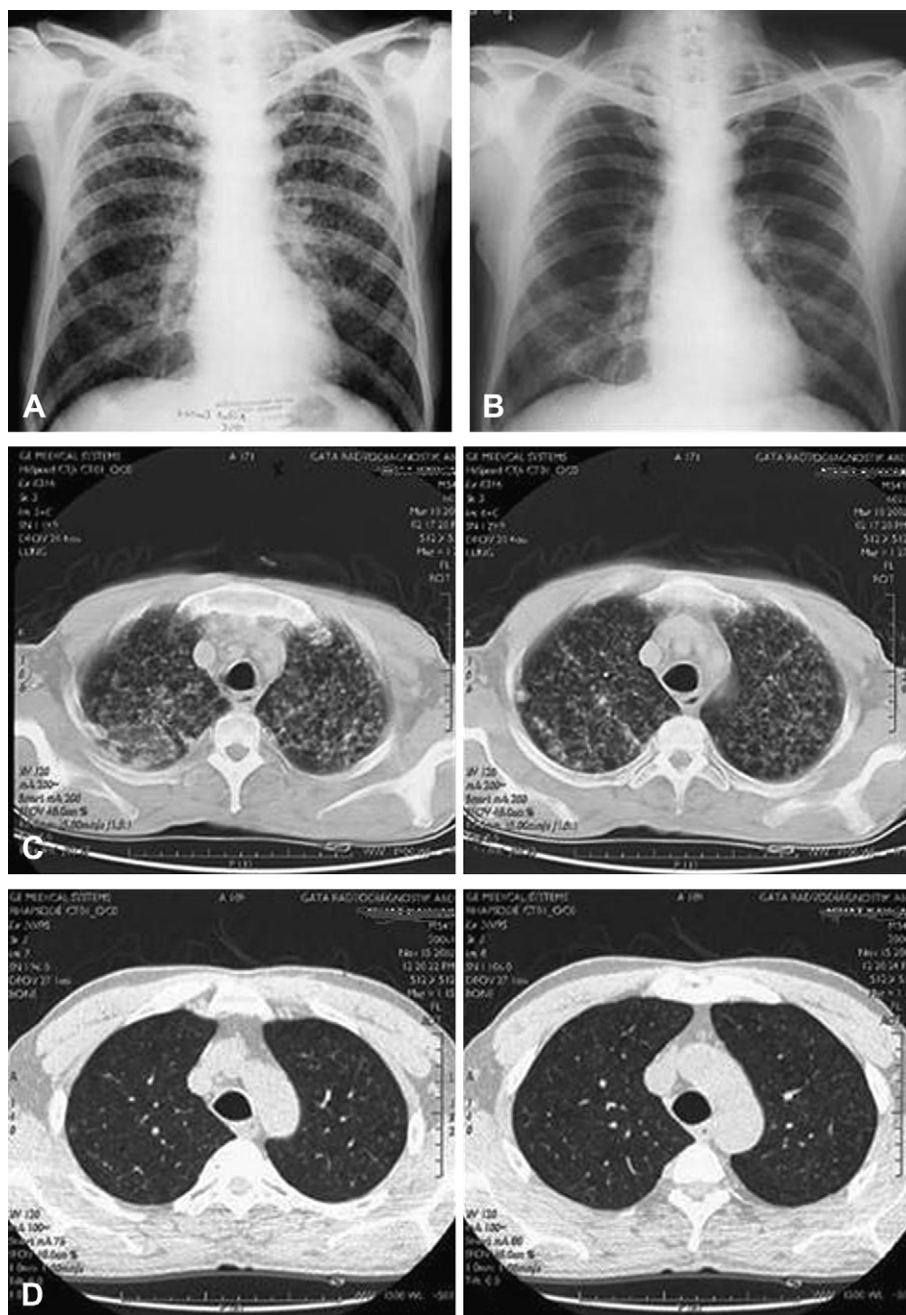


Figure 1 Initial posterior-anterior chest radiography showing bilaterally diffuse reticulo-nodular infiltration (A). Becoming almost normal following a 6-month-anti-tuberculous therapy (B). Initial thorax CT scan image showing bilaterally reticulo-nodular miliary infiltration (C). Almost all infiltrations being cleared at the end of anti-tuberculous therapy (D).

lymphadenopathies. Chest CT illustrated bilaterally diffuse reticulo-nodular pattern (Fig. 1).

After consulting the patient, with the department of urology, left orchidectomy was performed for histopathological diagnosis of the testicular mass. Histologic examination of the orchidectomy specimen revealed extended necrosis with caseous granulomatous inflammation of possible tuberculous etiology. Ziehl–Neelsen staining for acid-fast bacilli was positive.

An anti-tuberculous therapy, with isoniazid, rifampicin, ethambutol supplemented by pyrazinamide for the first 2 months, was started on the basis of pathologic findings. The patient became symptom free in the second month of the treatment. The abnormalities observed in chest radiography and in high resolution CT were almost completely resolved after 6 months (Fig. 1). Thus, the response to the treatment also confirmed the diagnosis as disseminated tuberculosis. Tumour markers decreased to the normal ranges at the end of the therapy as well (Table 1).

Discussion

Tuberculosis is the leading cause of death due to a single infectious agent worldwide.¹ Although the term “miliary TB” has been traditionally used for a pathological and radiological description, currently all forms of progressive widely disseminated haematogenous TB are being called as miliary TB, even the classical pathological or radiological findings are absent.² Many adult patients with progressive tuberculosis have accompanying medical conditions that cause their specific immune responses to wane. Our patient was not immunocompromised and he had no high risk condition associated with progressive tuberculosis.

Generally, when *M. tuberculosis* is identified in any specimen, other systems of patients with suspected tuberculosis are not tested. As a result of this inclination, involvement of multiple organs with TB is probably much more common than it is recognized; therefore, most of them are overlooked.² Abdominal involvement is a rare form of TB. CA-125 may be used as a marker for diagnosis, and in the follow-up of the patients with abdominal tuberculosis.^{3–6} CA-19-9 is expressed in mucous cells of the bronchial gland and surface of the bronchiolar surface epithelium cells in benign pulmonary diseases.⁷ It has been reported in several case reports that the level of CA-19-9 increased in various body fluids in patients with tuberculosis and decreased after anti-tuberculosis therapy. The authors suggested that CA-19-9 level may reflect the activity of pulmonary tuberculosis.^{7–9} CA-125 and CA-19-9 levels were found to be increased in our case. Plasma levels of these markers were started to decline after 2 months, and found within the normal levels at the end of anti-tuberculous therapy (Table 1). AFP level was only marginally elevated to 9.6 ng/mL and then decreased to

7.6 ng/mL over a six-month period. Therefore, it may not reflect the disease activity or response to therapy in tuberculosis. It has been known that tumour markers significantly decrease following orchidectomy in testicular tumour cases. In our case the elevation of tumour markers was considered as a reflection of disseminated tuberculosis, so tumour markers did not decreased immediately after orchidectomy.

Existence of a testicular mass was a coincidence that helped us to make the accurate diagnosis by an orchidectomy. It is not clear whether the actual cause of elevated levels of tumour markers is either abdominal or testicular involvement. To our knowledge this is the first in the literature where both tumour markers have been investigated simultaneously and found to be elevated in a patient with disseminated TB. We suggest that measurement of these markers may achieve a useful non-invasive tool in the diagnosis of disseminated tuberculosis. It may also be a useful to screen the response to anti-tuberculous therapy. However, it is sure that further studies in prospective design are needed to claim this hypothesis.

Conflict of interest statement

There is no conflict of interest for any of the authors of this case report.

References

1. Dolin PJ, Raviglione M, Kochi A. Global TB incidence and mortality during 1990–2000. *Bull World Health Organ* 1994;**72**: 213–20.
2. American Thoracic Society. Diagnostic standards and classification of tuberculosis in adults and children 1999. *Am J Respir Crit Care Med* 2000;**161**:1376–95.
3. Yilmaz A, Ece F, Bayramgürler B. The value of Ca 125 in the evaluation of tuberculosis activity. *Respir Med* 2001;**95**:666–9.
4. Mas MR, Comert B, Sağlamkaya U, et al. CA-125; a new marker for diagnosis and follow-up of patients with tuberculous peritonitis. *Dig Liver Dis* 2000;**3**:595–7.
5. Niwa Y, Kishimoto H, Shimokata K. Carcinomatous and tuberculous pleural effusions. Comparison of tumour markers. *Chest* 1985;**87**:351–5.
6. Simsek H, Savas MC, Kadayifci A, Tatar G. Elevated serum CA-125 concentration in patients with tuberculous peritonitis: a case control study. *Am J Gastroenterol* 1997;**92**:1174.
7. Takayama S, Kataoka N, Usui Y, et al. CA19-19 in patients with benign pulmonary diseases. *Nihon Kyobu Shikkan Gakkai Zasshi* 1990;**28**(10):1326–31.
8. Ishiura Y, Fujimura M, Minami S, et al. Increased CA19-19 level in serum and bronchoalveolar lavage fluid from a patient with pulmonary tuberculosis. *Nihon Kyobu Shikkan Gakkai Zasshi* 1996;**34**(4):477–81.
9. Komiya T, Matsushima T, Kimura M, Adachi M. A case of endo-bronchial tuberculosis with high serum CA19-19 and SLX level. *Kekkaku* 1994;**69**(10):615–9.